

**METHODS AND COMPOUNDS FOR MODULATING NUCLEAR RECEPTOR  
COACTIVATOR BINDING**

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**ABSTRACT OF THE DISCLOSURE**

The present invention relates to methods and agonist/antagonist compounds for modulating  
15 nuclear receptor coactivator binding. The invention includes a method for identifying residues  
comprising a coactivator binding site for a nuclear receptor of interest. Also included is a method  
of identifying agonists and/or antagonists that bind to a coactivator binding site of a nuclear  
receptor of interest. Agonists and antagonists of coactivator binding to nuclear receptors also are  
provided. The invention is exemplified by identification and manipulation of the coactivator  
20 binding site of the thyroid receptor (TR), and compounds that bind to this sites. The methods can  
be applied to other nuclear receptors including RAR, RXR, PPAR, VDR, ER, GR, PR, MR, and  
AR.